

## CLAIMS

1. A method of treatment of a patient undergoing opioid analgesic therapy which comprises minimising or mitigating the side effects of the opioid by the administration of a therapeutically effective amount of devazepide.
2. A method of treatment of a patient requiring analgesia which comprises the administration of a therapeutically effective amount of an opioid analgesic whilst minimising the side effects of the opioid by the separate, simultaneous or sequential administration of a therapeutically effective amount of devazepide.
3. A method according to claim 1 characterised in that the opioid is selected from the group morphine, or a salt thereof such as the sulphate, chloride or hydrochloride, or the other 1,4-hydroxymorphinan opioid analgesics such as naloxone, meperidine, butorphanol or pentazocine, or morphine-6-glucuronide, codeine, dihydrocodeine, diamorphine, dextropropoxyphene, pethidine, fentanyl, alfentanil, alphaprodine, buprenorphine, dextromoramide, diphenoxylate, dipipanone, heroin (diacetylmorphine), hydrocodone (dihydrocodeinone), hydromorphone (dihydromorphinone), levorphanol, meptazinol, methadone, metopon (methyldihydromorphinone), nalbuphine, oxycodone (dihydrohydroxycodeinone), oxymorphone (dihydrohydroxymorphinone), phenadoxone, phenazocine, remifentanyl, tramadol, or a salt of any of these.
4. A method according to claim 2 characterised in that the opioid is selected from the group morphine, or a salt thereof such as the sulphate, chloride or hydrochloride, or the other 1,4-hydroxymorphinan opioid analgesics such as naloxone, meperidine, butorphanol or pentazocine, or morphine-6-glucuronide, codeine, dihydrocodeine, diamorphine, dextropropoxyphene, pethidine, fentanyl, alfentanil, alphaprodine, buprenorphine, dextromoramide, diphenoxylate, dipipanone, heroin (diacetylmorphine), hydrocodone (dihydrocodeinone), hydromorphone (dihydromorphinone), levorphanol, meptazinol, methadone, metopon (methyldihydromorphinone), nalbuphine, oxycodone

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(dihydrohydroxycodone), oxymorphone (dihydrohydroxymorphine), phenadoxone, phenazocine, remifentanyl, tramadol, or a salt of any of these

5. A method according to claim 3 characterised in that the opioid is selected from the group hydromorphone, oxycodone, morphine and fentanyl.

6. A method according to claim 4 characterised in that the opioid is selected from the group hydromorphone, oxycodone, morphine and fentanyl.

10 7. A method according to claim 5 characterised in that the opioid is selected from the group morphine and morphine sulphate.

8. A method according to claim 6 characterised in that the opioid is selected from the group morphine and morphine sulphate.

15 9. A method according to claim 1 characterised in that the method of delivery of the devazepide and/or the opioid is selected from the group, administration intravenously, orally, intrathecally, intranasally, intrarectally, intramuscularly/subcutaneously, by inhalation and by transdermal patch.

20 10. A method according to claim 2 characterised in that the method of delivery of the devazepide and/or the opioid is selected from the group, administration intravenously, orally, intrathecally, intranasally, intrarectally, intramuscularly/subcutaneously, by inhalation and by transdermal patch.

25 11. A method according to claim 9 characterised in that the devazepide is administered intravenously or orally.

30 12. A method according to claim 10 characterised in that the devazepide is administered intravenously or orally.

13. A method according to claim 11 characterised in that the devazepide is administered orally.

5 14. A method according to claim 12 characterised in that the devazepide is administered orally.

10 15. A method according to claim 9 characterised in that the opioid is administered intravenously and the devazepide is administered intravenously.

16. A method according to claim 10 characterised in that the opioid is administered intravenously and the devazepide is administered intravenously.

15 17. A method according to claim 9 characterised in that the opioid is administered orally and the devazepide is administered orally.

18. A method according to claim 10 characterised in that the opioid is administered orally and the devazepide is administered orally.

20 19. A method according to claim 9 characterised in that the opioid is administered by intravenous administration or oral administration.

20. A method according to claim 10 characterised in that the opioid is administered by intravenous administration or oral administration.

25 21. A method according to claim 1 characterised in that the daily dosage of devazepide is up to 0.7 mg/kg/day.

30 22. A method according to claim 2 characterised in that the daily dosage of devazepide is up to 0.7 mg/kg/day.

23. A method according to claims 21 characterised in that the daily dosage of devazepide is from 25  $\mu\text{g/kg/day}$  to 0.7  $\text{mg/kg/day}$ .

5 24. A method according to claims 22 characterised in that the daily dosage of devazepide is from 25  $\mu\text{g/kg/day}$  to 0.7  $\text{mg/kg/day}$ .

25. A method according to claim 23 characterised in that the daily dosage of devazepide is from 50  $\mu\text{g/kg/day}$  to 0.5  $\text{mg/kg/day}$ .

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26. A method according to claim 24 characterised in that the daily dosage of devazepide is from 50  $\mu\text{g/kg/day}$  to 0.5  $\text{mg/kg/day}$ .

15 27. A method according to claim 25 characterised in that the devazepide is administered orally and the daily dosage of devazepide is from 0.07  $\text{mg/kg/day}$  to 0.29  $\text{mg/kg/day}$ .

20 28. A method according to claim 26 characterised in that the devazepide is administered orally and the daily dosage of devazepide is from 0.07  $\text{mg/kg/day}$  to 0.29  $\text{mg/kg/day}$ .

25 29. A method according to either of claims 25 characterised in that the devazepide is administered intravenously administration the dosage of devazepide is from 50  $\mu\text{g/kg/day}$  to 0.5  $\text{mg/kg/day}$ .

30. A method according to either of claims 26 characterised in that the devazepide is administered intravenously administration the dosage of devazepide is from 50  $\mu\text{g/kg/day}$  to 0.5  $\text{mg/kg/day}$ .

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31. A method according to claim 1 characterised in that the daily dosage of the opioid is from 5 to 2000mg daily.

5 32. A method according to claim 2 characterised in that the daily dosage of the opioid is from 5 to 2000mg daily.

33. A method according to claim 31 characterised in that the daily dosage of the opioid is from 5 to 100mg daily.

10 34. A method according to claim 32 characterised in that the daily dosage of the opioid is from 5 to 100mg daily.

15 35. A method according to claim 1 characterised in that the side effect which is inhibited, mitigated or minimised is selected from the group, constipation, dizziness, tiredness/fatigue and vomiting.

20 36. A method according to claim 2 characterised in that the side effect which is inhibited, mitigated or minimised is selected from the group, constipation, dizziness, tiredness/fatigue and vomiting.

37. A method according to claim 1 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

25 38. A method according to claim 2 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

39. The use of devazepide in the manufacture of a medicament which inhibits or mitigates the undesirable side effects of administration of a therapeutically effective amount of an opioid analgesic.

40. The use according to claim 39 characterised in that the devazepide used is the S enantiomer wherein the level of R enantiomer is not greater than 1.5% w/w.

5 41. The use of devazepide in the manufacture of a medicament for use in the method of either of claim 1.

42. The use of devazepide in the manufacture of a medicament for use in the method of either of claim 2.

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